

- 1 2. (As filed) The method of claim 1, wherein angiogenesis activity is decreased.
- 1 3. (As filed) The method of claim 1, wherein angiogenesis activity is increased.
- 1 4. (As filed) The method of claim 2, wherein the response is a decrease in normal blood  
2 vessel formation.
- 1 5. (As filed) The method of claim 3, wherein the response is an increase in normal blood  
2 vessel formation.
- 1 6. (As filed) The method of claim 2, wherein the response is loss of existing blood  
2 vessels.
- 1 7. (As filed) The method of claim 1, wherein the teleost is an embryo, larva, or adult.
- 1 8. (As filed) The method of claim 1, wherein the teleost is a zebrafish, medaka, Giant  
2 rerio, or puffer fish.
- 1 9. (As filed) The method of claim 8, wherein the teleost is a zebrafish embryo.
- 1 10. (As filed) The method of claim 1, wherein the teleost is a wildtype strain.
- 1 11. (As filed) The method of claim 1, wherein the teleost contains a mutation in a selected  
2 gene.
- 1 12. (As filed) The method of claim 1, wherein the teleost is transgenic.
- 1 13. (As filed) The method of claim 1, wherein the agent is administered to the teleost by  
2 dissolving the agent in media containing the teleost.
- 1 14. (As filed) The method of claim 1, wherein the agent is administered to the teleost by  
2 injecting the agent into the teleost.
- 1 15. (As filed) The method of claim 1, wherein the agent is administered to the teleost in  
2 conjunction with a carrier.
- 1 16. (As filed) The method of claim 15, wherein the carrier is a solvent, lipid, or peptide.
- 1 17. (As filed) The method of claim 1, wherein the agent is a compound and a library of  
2 compounds is screened for angiogenesis activity.

1                   18.     (As filed) The method of claim 1, wherein the agent is a nucleic acid, peptide, protein,  
2 glycoprotein, carbohydrate, lipid, or glycolipid.

1                   19.     (As filed) The method of claim 18, wherein the nucleic acid is DNA or RNA.

1                   20.     (As filed) The method of claim 5, wherein blood vessels are visualized by light  
2 microscopy after alkaline phosphatase staining of the teleost.

1                   21.     (Amended) A method of screening an agent for an effect on cell death  
2 activity, said method comprising contacting a living teleost post 12-hours of development with  
3 a dye with affinity for dead cells, and thereafter administering the agent to be screened to [a]  
4 the living teleost and detecting [a response] the dye in at least one specific tissue or organ in  
5 the living teleost indicating an effect on cell death activity in at least one specific tissue or  
6 organ of the living teleost.

1                   43.     (Amended) A method of screening an agent for toxic activity in vivo  
2 comprising administering the agent to a teleost in vivo and detecting a change in level of an  
3 enzyme or mRNA in at least one tissue or organ of the teleost responsive to the agent [response  
4 in the teleost] indicating toxic activity in the at least one tissue or organ of the teleost.

Please add the following new claims:

1                   --54.   (New) The method of claim 21, wherein the response is an increase in  
2 cell death activity.

1                   55.     (New) The method of claim 21, wherein the response is a decrease in  
2 cell death activity.

1                   56.     (New) The method of claim 21, wherein the response is an increase in  
2 apoptotic activity or necrotic activity.

1                   57.     (New) The method of claim 21, wherein the response is a decrease in  
2 apoptotic activity or necrotic activity.

1                   58.     (New) The method of claim 56, wherein the increase in apoptotic  
2 activity comprises an increase in cell death in a tissue or organ of the teleost.

1                   59.     (New) The method of claim 57, wherein the decrease in apoptotic  
2 activity comprises a decrease in cell death in a tissue or organ of the teleost.

1                   60.     (New) The method of claim 54, wherein the method further comprises  
2 detecting a response in cell death activity in the teleost after a predetermined period of time,  
3 said time being sufficient for detectable differences in cell death activity to occur in the  
4 presence of the agent.

1                   61.     (New) The method of claim 56, wherein the increase in apoptotic  
2 activity is detected by light microscopy or fluorescence microscopy.

1                   62.     (New) The method of claim 21, wherein the agent is administered to the  
2 teleost by dissolving the agent in media containing the teleost.

1                   63.     (New) The method of claim 21, wherein a fluorescent dye which labels  
2 dead or dying cells is administered to the teleost prior to administration of the agent to the  
3 teleost.

1                   64.     (New) The method of claim 63, wherein the fluorescent dye is  
2 administered to the teleost by dissolving the fluorescent dye in media containing the teleost.

1                   65.     (New) The method of claim 63, wherein the fluorescent dye is  
2 administered to the teleost by injecting the fluorescent dye into the teleost.

1                   66.     (New) The method of claim 63, further comprising administering the  
2 agent to the teleost by dissolving the agent in the media containing the teleost or injecting the  
3 agent into the teleost after administration of the fluorescent dye to the teleost.

1                   67.     (New) The method of claim 63, wherein a fluorescent dye is a  
2 monomeric cyanine dye.

1                   68.   (New) The method of claim 67, wherein the fluorescent dye is  
2 benzothiazolium-4-quinolium dye.

1                   69.   (New) The method of claim 21, wherein the teleost is a zebrafish.

1                   70.   (New) The method of claim 54, wherein the increase in cell death  
2 activity is detected in more than one tissue or organ of the teleost simultaneously.

1                   71.   (New) The method of claim 70, wherein the increase in cell death  
2 activity is detected in more than one tissue or organ of the teleost simultaneously over time at  
3 predetermined intervals.

1                   72.   (New) The method of claim 60, wherein the method further comprises  
2 detecting the increase in cell death activity over time at predetermined intervals.

1                   73.   (New) The method of claim 56, wherein the increase in apoptotic  
2 activity or necrotic activity is detected in at least one organ or tissue or combination thereof.

1                   74.   (New) The method of claim 21, wherein the agent is a compound and a  
2 library of compounds is screened for an effect on cell death activity.

1                   75.   (New) The method of claim 43, wherein the response in the teleost  
2 indicating toxic activity is detected over time.

1                   76.   (New) The method of claim 43, wherein the response in the teleost  
2 indicating toxic activity is detected in at least two tissues, at least two organs, or at least one  
3 tissue and one organ simultaneously.

1                   77.   (New) The method of claim 76, wherein the response in the teleost  
2 indicating toxic activity is over time at predetermined intervals.

1                   78.   (New) The method of claim 43, further comprising administering the  
2 agent to at least two teleosts and detecting a response indicating toxic activity in each of said at  
3 least two teleosts simultaneously.

1                   79.     (New) The method of claim 78, wherein each of said at least two  
2 teleosts is contained in a well of a multi-well plate.

1                   80.     (New) The method of claim 1, further comprising screening the agent  
2 for toxic activity by detecting a response in the teleost indicating toxic activity.

1                   81.     (New) The method of claim 1, further comprising screening the agent  
2 for an ability to enhance or inhibit cell death activity by detecting a response in the teleost  
3 indicating an enhancement or inhibition of cell death activity.

1                   82.     (New) The method of claim 21, further comprising screening the agent  
2 for toxic activity by detecting a response in the teleost indicating toxic activity.

1                   83.     (New) The method of claim 1, wherein the teleost is bleached after  
2 staining with alkaline phosphatase.

1                   84.     (New) The method of claim 21, wherein the method is conducted in a  
2 teleost *in vivo*.

1                   85.     (New) The method of claim 1, wherein the teleost is contained in a  
2 microtiter well.

1                   86.     (New) The method of claim 85, wherein the response is detected using a  
2 microplate reader.

1                   87.     (New) The method of claim 21, wherein the teleost is contained in a  
2 microtiter well.

1                   88.     (New) The method of claim 87, wherein the dye in at least one specific  
2 tissue or organ is detected using a microplate reader.

1                   89.     (New) The method of claim 43, wherein the teleost is contained in a  
2 microtiter well.